

-continued

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<223> OTHER INFORMATION: Synthetic- Murine leukemia virus epitope used
in H-2Ld MuLV gp70 Tetramer

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<223> OTHER INFORMATION: Synthetic- Influenza A virus epitope used in
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What is claimed is:

1. Modified tumor-derived extracellular vesicles (EVs) isolated from a tumor cell having reduced or lacking expression of an immune suppressive factor.

2. The modified tumor-derived EVs of claim 1, wherein the immune suppressive factor is miR-424.

3. The modified tumor-derived EVs of claim 1, wherein the EVs are exosomes or microvesicles.

4. The modified tumor-derived EVs of claim 1, wherein the EVs are about 30 to about 300 nanometers in diameter.

5. The modified tumor-derived EVs of claim 1, wherein the EVs are isolated from a tumor cell that has been modified to inhibit expression of immune suppressive factor.

6. The modified tumor-derived EVs of claim 1, wherein the tumor cell is a cultured tumor cell or a tumor organoid.

7. The modified tumor-derived EVs of claim 6, wherein the tumor cell is selected from the group consisting of a colorectal cancer cell, breast cancer cell, endometrial cancer cell, prostate cancer cell, lung cancer cell, melanoma cell, and pancreatic cancer cell.

8. The modified tumor-derived EVs of claim 1, wherein the EVs comprise one or more exogenous antigens.

9. A composition comprising the modified tumor-derived EVs of claim 1, a pharmaceutically acceptable carrier, and one or more checkpoint inhibitor.

10. A method of producing a tumor-derived extracellular vesicle (EV) substantially lacking expression of an immune suppressive factor, the method comprising:

(a) providing a modified tumor cell that lacks expression of the immune suppressive factor and can produce EVs; and

(b) isolating EVs produced by the tumor cell, wherein the EVs substantially lack expression of the immune suppressive factor.

11. The method of claim 10, wherein the immune suppressive factor is miR-424.

12. The method of claim 10, wherein the method comprises:

a) transducing the tumor cell with an anti-miR-424 expression vector,

b) transfecting the tumor cells with an anti-miRNA-424 oligo, or

c) knocking out the miR-424 gene from the tumor cell using CRISP/Cas9 genome editing.

13. The method of any one of claim 10, wherein the tumor cell is a cultured tumor cell or a tumor organoid.

14. A method of treating cancer in a subject, the method comprising: administering an effective amount of tumor-derived EVs to a subject in need thereof, wherein the EVs substantially lack expression of an immune suppressive factor or miR-424, and wherein the EVs comprise one or more cancer antigen.

15. The method of claim 14 further comprising administering a chemotherapy to the subject.

16. The method of claim 15, wherein the tumor-derived EVs are administered after the subject's T-cell population have substantially recovered following the chemotherapy.

17. A method of stimulating an anti-tumor response in a subject having cancer, the method comprising: administering a vaccine comprising the tumor-derived EVs of claim 1 to a subject having cancer in an amount effective to elicit an anti-tumor response in the subject.

18. The method of claim 17, wherein administering the vaccine or composition (a) increases CD28 expression on T cells, (b) increases T cell proliferation, or (c) both increases CD28 expression on T cells and increases T cell proliferation in a subject having cancer.

19. The method of claim 17, wherein the anti-tumor response comprises the reduction or inhibition of secondary tumor growth.

20. A method of sensitizing a tumor cell in a subject to immune checkpoint inhibitors, the method comprising: administering the tumor-derived EVs of claim 1 to a subject having cancer and subsequently administering one or more checkpoint inhibitor to the subject.

21. The method of claim 20, wherein the checkpoint inhibitor is selected from a PD-1 inhibitor, a PDL1 inhibitor, a CTL4 inhibitor, or a combination thereof.

22. A method of preventing, reducing, or inhibiting tumor growth in a subject, the method comprising: administering an effective amount of a vaccine comprising tumor-derived EVs of claim 1 to a subject in need thereof to prevent, reduce